U.S. PATENT DOCUMENTS						
EXAMINER INITIAL	DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE (IF APPROPRIATE)
AU /M 1	6,239,171	May 29, 2001	Lane et al.			

FOREIGN PATENT DOCUMENTS							
EXAMINER		DATE	COUNTRY	CLASS	SUBCLASS	TRANSLATION	
INITIAL	DOCUMENT NUMBER	UATE	COUNTY	00.00	00000	YES	NO
							_
-							

EXAMINER INITIAL		OTHER DOCUMENTS (INCLUDING AUTHOR, TITLE, DATE, PERTINENT PAGES, ETC.)
MM	2	Sardi, B., Special Report: Part II Choosing Natural Agents for Cholkesterol Control, March 2003.
	3	
	4	
	5	
	6	
	7	
	8	
	9	
	10	
	11	
	12	
	13	·
	14	
	15	
	16	
	17	
	18	

	01.1 [1					/	
	AM/A		/	4	4	91	
EXAMINER		DATE CONSIDERED	·/ ($/\!\!\!/$		7/0	

*EXAMINER: INITIAL IF CITATION CONSIDERED, WHETHER OR NOT CITATION IS IN CONFORMANCE WITH MPEP 609, DRAW LINE THROUGH CITATION IF NOT IN CONFORMANCE AND NOT CONSIDERED, INCLUDE COPY OF THIS FORM WITH NEXT COMMUNICATION TO APPLICANT.

SH	FFT	3 C	IF 6

FORM PTO 449 U.S. DEPARTMENT OF COMMERCE ATTY. DOCKET NO. BT-002

PADE PATENT AND TRADEMARK OFFICE

INFORMATION DISCLOSURE STATEMENT
BY APPLICANT
BY APPLICANT

(USE SEVERAL SHEETS IF NECESSARY)

ATTY. DOCKET NO. 8T-002	APPLICATION NO. 10/821,679
APPLICANT Tan, Barrie et al.	
FILING DATE April 8, 2004	GROUP Unknown

U.S. PATENT DOCUMENTS								
EXAMII INITI			DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE (IF APPROPRIATE)
W	M	1	2003139467	July 24, 2003	Igarashi			•
	1	2	6441029	August 27, 2002	Elson			
		3	6358997	March 19, 2002	Clark			-
		4	6350453	February 26, 2002	Tan			
		5	6083979	July 4, 2000	Sehti			
		6	5756109	May 26, 1998	Burger			
		7	5660691	August 26, 1997	Barnicki			
		8	5602184	February 11, 1997	Myers			
		9	5318993	June 7, 1994	Pearce			
Α		16	5217992	June8, 1993	Wright			1
		11	5157132	October 20, 1992	Tan			
T	1							

FOREIGN PATENT DOCUMENTS							
EXAMINER	DOCUMENT NUMBER DATE COUNTRY CLASS	SUBCLASS	TRANSLATION				
INITIAL .	JOGOMENT NOMBEN	UNIC				YES	NO
			· · · · · · · · · · · · · · · · · · ·				

EXAMINER OTHER DOCUMENTS (INCLUDING AUTHOR, TITLE, DATE, PERTINENT PAGES	, ETC.)
Adachi, J. and G. Ioannidis (2000). "Glucocorticoid-induced osteoporos Research 49: 120-134.	sis." Drug Development
Amin, D., R. Rutledge, et al. (1997). "RPR 107393, a potent squalene s orally effective cholesterol-lowering agent: comparison with inhibitors J Pharmacol Exp Ther. 281(2): 746-752.	
Ayanian, J., C. Fuchs, et al. (1988). "Lovastatin and rhabdomyolysis." A	Ann Intern Med. 109: 682-
Baker, S. and M. Tarnopolsky (2001). "Statin myopathies: pathophysio perspectives." Clin Invest Med. 24(5): 258-272.	logic and clinical

EXAMINER DATE CONSIDERED

*EXAMINER: INITIAL IF CHATION CONSIDERED, WHETHER OR NOT CITATION IS IN CONFORMANCE WITH MPEP 809; DRAW LINE THROUGH CITATION IF NOT IN CONFORMANCE AND NOT CONSIDERED, INCLUDE COPY OF THIS FORM WITH NEXT COMMUNICATION TO APPLICANT.

		STILET Z OF
FORM PTO-1449 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE	ATTY, DOCKET NO. BT-002	APPLICATION NO. 10/821,679
INFORMATION DISCLOSURE STATEMENT BY APPLICANT	APPLICANT Ten, Barrie et al.	
(USE SEVERAL SHEETS IF NECESSARY)	FILING DATE April 8, 2004	GROUP Unknown

EXAMINER INITIAL		OTHER DOCUMENTS (INCLUDING AUTHOR, TITLE, DATE, PERTINENT PAGES, ETC.)
WN	16	Ballantyne, C., A. Corsini, et al. (2003). "Risk for myopathy with statin therapy in high-risk patients." Arch. Intern. Med. 163(5): 553-564.
	17	Bentinger, M., J. Grunler, et al. (1998). "Phosphorylation of farnesol in rat liver microsomes: properties of farnesol kinase and farnesyl phosphate kinase." Arch Biochem Biophys 353(2): 191-198.
	18	Bliznakov, E. (2002). "Lipid-lowering drugs (statins), cholesterol, and coenzyme Q10. The Baycol casea modern Pandora's box." Biomed Pharmacother. 56(1): 56-59.
	19	Cenedella, R. (1995). "Role of transcription, translation, and protein turnover in controlling the distribution of 3-hydroxy-3-methylglutaryl coenzyme A reductase in the lens." Invest Ophthalmol Vis Sci. 36(10): 2133-2141.
	20	Cenedella, R. (1997). "Posttranscriptional regulation of 3-hydroxy-3-methylglutaryl coenzyme A reductase in lens epithelial cells by mevalonate-derived nonsterols." Exp Eye Res. 65(1): 63-72.
	21	Chamberlain, L. (2001). "Inhibition of isoprenoid biosynthesis causes insulin resistance in 3T3-L1 adipocytes." FEBS Letters 507(3): 357-361.
	22	Ciosek, C., D. Magnin, et al. (1993). "Lipophilic 1,1-bisphosphonates are potent squalene synthase inhibitors and orally active cholesterol lowering agents in vivo." J. Biol. Chem 268: 24832-24837.
	23	Colli, S., S. Eligini, et al. (1997). "Vastatins inhibit tissue factor in cultured human macrophages. A novel mechanism of protection against atherothrombosis." Arterioscler Thromb Vasc Biol 17: 265-272.
	24	Craveiro, A. A., et al., (1989). "The Presence of Geranylgeraniol in Bixa Orelana Linn.," Quimica Nova, 12(3):297-298.
	25	Cruz, A. and B. Gruber (2002). "Statins and osteoporosis: Can these lipid-lowering drugs also bolster bones?" Cleveland Clinic Journal of Medicine 69(4): 277-288.
	26	Danesi, R., C. McLellan, et al. (1995). "Specific labeling of isoprenylated proteins: application to study inhibitors of the post-translational farnesylation and geranylgeranylation." Biochem Biophys Res Commun. 206(2): 637-643.
	27	DeFronzo, R. A. (1998). "Pathogenesis of Type 2 Diabetes: Metabolic & Molecular Implications for Identifying Diabetes Genes." Ann. Rev. Diabetes: 1-93.
	28	Flint, O., B. Masters, et al. (1997). "HMG CoA reductase inhibitor-induced myotoxicity: pravastatin and lovastatin inhibit the geranylgeranylation of low-molecular-weight proteins in neonatal rat muscle cell culture." Toxicol Appl Pharmacol. 145(1): 99-110.
	29	Flint, O., B. Masters, et al. (1997). "Inhibition of cholesterol synthesis by squalene synthase inhibitors does not induce myotoxicity in vitro." Toxicol Appl Pharmacol 145(1): 91-98.
	30	Folkers, K., A. Osterborg, et al. (1997). "Activities of vitamin Q10 in animal models and a serious deficiency in patients with cancer." Biochem Biophys Res Commun. 234(2): 296-299.

EXAMINER DATE CONSIDERED

*EXAMINER: INITIAL IF CITATION CONSIDERED, WHETHER OR NOT CITATION IS IN CONFORMANCE WITH MPEP 609; DRAW LINE THROUGH CITATION IF NOT IN CONFORMANCE AND NOT CONSIDERED, INCLUDE COPY OF THIS FORM WITH NEXT COMMUNICATION TO APPLICANT.

FORM PTO-1449 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE	ATTY. DOCKET NO. BT-002	APPLICATION NO. 10/821,679	
INFORMATION DISCLOSURE STATEMENT BY APPLICANT	APPLICANT Tan, Barrie et al.		
(USE SEVERAL SHEETS IF NECESSARY)	FILING DATE April 8, 2004	GROUP Unknown	

EXAMINER INITIAL		OTHER DOCUMENTS (INCLUDING AUTHOR, TITLE, DATE, PERTINENT PAGES, ETC.)	
NWM	31	Gavish, D., E. Leibovitz, et al. (2000). "Bezafibrate and simvastatin combination therapy for diabetic dyslipidaemia: efficacy and safety." J. Intern. med. 247: 563-569.	
	Grünler, J., J. Ericsson, et al. (1994). "Branch-point reactions in the biosynthesis of cholesterol, dolichol, ubiquinone and prenylated proteins." Biochim Biophys Acta. 1212(3): 259-77.		
	Guillet-Deniau, I., et al. (2003). "Glucose induces de novo fatty acid synthesis in rat skeletal muscle through a SREBP-1c dependent pathway." Diabetes 52(Suppl. 1): extended abstract. 1024P.		
	34	Hamilton-Craig, I. (2001). "Statin-associated myopathy." Med J Aust. 175(9): 486-489.	
-	Haslinger, B., M. Goedde, et al. (2002). "Simvastatin Increases Fibrinolytic Activity in Human Peritoneal Mesothelial Cells Independent of Cholesterol Lowering." Kidney International 62(5): 1611-1619.		
	Hevener, A., W. He, et al. (2003). "Muscle-specific Pparg deletion causes insulin resistance." Med. 9(12): 1491-1497.		
	Holdaas, H., A. Jardine, et al. (2001). "Effect of fluvastatin on acute renal allograft rejection: a randomized multicenter trial." Kidney Int. 60(5): 1990-1997.		
	Johnston, S. and L. Kelland (2001). "Farnesyl transferase inhibitorsa novel therapy for breast cancer." Endocr Relat Cancer. 8(3): 227-235.		
$\lceil \cdot \rceil$	Jondiko, I. J. O. and Pattenden, G., (1989). "Terpenoids and an Apocarotenoid from Seeds of Bixa Orellana," Phytochemistry, 28(11):3159-3162.		
	Judy, W., T. Nguyen, et al. (2004). "Evidence for regression of PSA and tumor size in patients with prostate cancer." Submitted for publication in Proc. Natl. Acad. Sci.		
	41 Kraegen, E. (1998). "Physiologic manifestations of PPAR-gamma activation: preclinical stud Clinical Courier 16(48): 5-7.		
	42	Lankin, V., A. Tikhaze, et al. (2000). "Intensification in vivo of free radical oxidation of low density lipoproteins in plasma from patients with myocardial ischemia treated by HMG-CoA-reductase pravastatin and suppression of lipid peroxidation by ubiquinone Q10." Biull Eksp Biol Med. 129(2): 176-179.	
	43	Lehmann, J., L. Moore, et al. (1995). "An antidiabetic thiazolidinedione is a high affinity ligand for peroxisome proliferator-activated receptor gamma (PPAR gamma)." J Biol Chem. 270(22): 12953-12956.	
	44	Luckman, S., F. Coxon, et al. (1998). "Heterocycle-containing bisphosphonates cause apoptosis and inhibit bone resorption by preventing protein prenylation: evidence from structure-activity relationships in J774 macrophages." J Bone Miner Res. 13(11): 1668-1678.	
	45	McCarty, M. (1999). "Can correction of sub-optimal coenzyme Q status improve beta-cell function in type II diabetics?" Med Hypotheses. 52(2): 397-400.	

EXAMINER DATE CONSIDERED DATE CONSIDERED

*EXAMINER: INITIAL IF CITATION CONSIDERED, WHETHER OR NOT CITATION IS IN CONFORMANCE WITH MPEP 609; DRAW LINE THROUGH CITATION IF NOT IN CONFORMANCE AND NOT CONSIDERED, INCLUDE COPY OF THIS FORM WITH NEXT COMMUNICATION TO APPLICANT.

FORM PTO-1449	U.S. DEPARTMENT OF COMMERCE
PATENT	AND TRADEMARK OFFICE

APPLICANT Tan, Barrie et al.

ATTY. DOCKET NO. BT-002

APPLICATION NO. 10/821,679

INFORMATION DISCLOSURE STATEMENT BY APPLICANT

(USE SEVERAL SHEETS IF NECESSARY)

FILING DATE April 8, 2004 GROUP Unknown

EXAMINER INITIAL			OTHER DOCUMENTS (INCLUDING AUTHOR, TITLE, DATE, PERTINENT PAGES, ETC.)	
$\int_{\mathcal{U}}$	MM	McGuire, T. and S. Sebti (1997). "Geranylgeraniol potentiates lovastatin inhibition of oncog H-Ras processing and signaling while preventing cytotoxicity." Oncogene 14: 305-312.		
*	47 McLaughlin, e. a. (2003). "Prediction of IR with plasma TG or TG/HDL ratio." Diab		McLaughlin, e. a. (2003). "Prediction of IR with plasma TG or TG/HDL ratio." Diabetes 52(Suppl. 1): A224 extended abstract. 962P.	
		Miller, P. (2001). "Bisphosphonates for the prevention and treatment of corticosteroid-induce osteoporosis." Osteoporos Int. Suppl 3: S3-S10.		
		49	Miquel, K., A. Pradines, et al. (1998). "Competitive Inhibition of Choline Phosphotransferase by Geranylgeraniol and Farnesol Inhibits Phosphatidylcholine Synthesis and Induces Apoptosis in Human Lung Adenocarcinoma A549 Cells." J. of Biol. Chem. 273(40): 26179-26186.	
		50	Olbricht, C., C. Wanner, et al. (1997). "Accumulation of lovastatin, but not pravastatin, in the blood of cyclosporine-treated kidney graft patients after multiple doses." Clin Pharmacol Ther. 62(3): 311-321.	
		Omar, M., J. Wilson, et al. (2001). "Rhabdomyolysis and HMG-CoA reductase inhibitors." The Annals of Pharmacotherapy 35(9): 1096-1107.		
		52	Ownby, S. and R. Hohl (2002). "Farnesol and geranylgeraniol: prevention and reversion of lovastatin-induced effects in NIH3T3 cells." Lipids 37(2): 185-92.	
П		53	Pearce, B., R. Parker, et al. (1992). "Hypocholesterolemic activity of synthetic and natural tocotrienols." J Med Chem. 35(20): 3595-3606.	
		54		
		Poels, P. and F. Gabreels (1993). "Rhabdomyolysis: a review of the literature." Clin. Neurol. Neurosurg. 95: 175-192.		
		Polverino, A. and S. Patterson (1997). "Selective activation of caspases during apoptotic induction in HL-60 cells. Effects Of a tetrapeptide inhibitor." J Biol Chem. 272(11): 7013-7021.		
П		57	Qureshi, et al., (1995). "Response of Hypercholesterolemic Subjects to Administration of Tocotrienols," Lipids, 30(12).	
		58	Raner, G., A. Muir, et al. (2002). "Farnesol as an inhibitor and substrate for rabbit liver microsomal P450 enzymes." Biochem Biophys Res Commun. 293(1): 1-6.	
		59	Regazzi, M., I. Iacona, et al. (1993). "Altered disposition of pravastatin following concomitant drug therapy with cyclosporin A in transplant recipients." Transplant Proc. 25: 2732-2734.	
		60	Rogers, M. (2000). "Statins: lower lipids and better bones?" Nature Medicine 6(1): 21-23.	
		61	Saito, J., J. Davis, et al. (1995). "Users of low-dose glucocorticoids have increased bone loss rates: a longitudinal study." Calcif Tissue Int. 57(2): 115-119.	
		62	Schlienger, R., W. Haefeli, et al. (2001). "Risk of cataract in patients treated with statins." Arch Intern Med. 161(16): 2021-2026.	

EXAMINER

DATE CONSIDERED

*EXAMINER: INITIAL IF CITATION CONSIDERED, WHETHER OR NOT CITATION IS IN CONFORMANCE WITH MPEP 609; DRAW LINE THROUGH CITATION IF NOT IN CONFORMANCE AND NOT CONSIDERED, INCLUDE COPY OF THIS FORM WITH NEXT COMMUNICATION TO APPLICANT.

	<u> </u>	
FORM PTO-1449 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE	ATTY. DOCKET NO. BT-002	APPLICATION NO. 10/821,679
INFORMATION DISCLOSURE STATEMENT BY APPLICANT	APPLICANT Tan, Barrie et al.	
(USE SEVERAL SHEETS IF NECESSARY)	FILING DATE April 8, 2004	GROUP Unknown

EXAMINER INITIAL		OTHER DOCUMENTS (INCLUDING AUTHOR, TITLE, DATE, PERTINENT PAGES, ETC.)	
MN	6 3	Smith, P., R. Eydelloth, et al. (1991). "HMG-CoA reductase inhibitor-induced myopathy in the rat: cyclosporine A interaction and mechanism studies." J Pharmacol Exp Ther. 257(3): 1225-1235.	
	64 -	Smith, S. (1998). "The molecular pharmacology of PPAR-gamma." Clinical Courier 16(48): 3-4.	
	Spach, D., J. Bauwens, et al. (1991). "Rhabdomyolysis associated with lovastatin and erythromycin use." West J. Med. 154: 213-215.		
Staels, B., J. Dallongeville, et al. (1998). "Mechanism of action of fibrates on lipid an lipoprotein metabolism." Circulation 98(19): 2088-2093.		lipoprotein metabolism." Circulation 98(19): 2088-2093.	
	67	Stark, W., M. Blaskovich, et al. (1998). "Inhibiting geranylgeranylation blocks growth and promotes apoptosis in pulmonary vascular smooth muscle cells." Am J Physiol. 275: L55-L63.	
		Takahashi, N., T. Kawada, et al. (2002). "Dual action of isoprenols from herbal medicines on both PPARgamma and PPARalpha in 3T3-L1 adipocytes and HepG2 hepatocytes." FEBS Lett. 514(2-3): 315-322.	
	69	Talley, N., A. Weaver, et al. (1992). "Onset and disappearance of gastrointestinal symptoms and functional gastrointestinal disorder." Am J Epidemiol. 136(2): 165-177.	
	70	Thai, L., J. S. Rush, et al. (1999). "Farnesol is utilized for isoprenoid biosynthesis in plant cells via farnesyl pyrophosphate formed by successive monophosphorylation reactions." Proc Natl Acad Sci 96(23): 13080-13085.	
	Thibault, A., D. Samid, et al. (1996). "Phase I study of lovastatin, an inhibitor of the mevalonate pathway, in patients with cancer." Clin Cancer Res. 2(3): 484-491.		
	72 Theriault, A., et al., (1999). "Tocotrienol: A Review of Its Therapeutic Potential," Clinical Biochemistry, 32(5):309-319.		
	73 VERIS (1989). The role of Vitamin E in exercise, Vitamin E Research & Information Serv (November).		
	74	VERIS (1990). Protective role of vitamin E in cataract development, Vitamin E Research & Information Service (February).	
	Watkins, T. R. and Bierenbaum, M. L., (1999). "Tocotrienols: Biological and Health Effe Antioxidant Status, Diet, Nutrition, and Health, Andreas M. Papas, ed. (CRC Press) pp. 4		
	76	Watts, G., C. Castelluccio, et al. (1993). "Plasma coenzyme Q (ubiquinone) concentrations in patients treated with simvastatin." J Clin Pathol. 46(11): 1055-1057.	
	77	Watts, N., D. Freedholm, et al. (1999). "The clinical tolerability profile of alendronate." Int J Clin Pract Suppl. 101: 51-61.	
	78	Willson, T., M. Lambert, et al. (2001). "Peroxisome proliferator-activated receptor gamma and metabolic disease." Annu Rev Biochem. 70: 341-367.	

EXAMINER DATE CONSIDERED

*EXAMINER: INITIAL IF CITATION CONSIDERED, WHETHER OR NOT CITATION IS IN CONFORMANCE WITH MPEP 609, DRAW LINE THROUGH CITATION IF NOT IN CONFORMANCE AND NOT CONSIDERED, INCLUDE COPY OF THIS FORM WITH NEXT COMMUNICATION TO APPLICANT.

SHEET 6 OF 6

FÖRM PTO-1449 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE	ATTY, DOCKET NO. BT-002	APPLICATION NO. 10/821,679
INFORMATION DISCLOSURE STATEMENT BY APPLICANT	APPLICANT Tan, Barrie et al.	
(USE SEVERAL SHEETS IF NECESSARY)	FILING DATE April 8, 2004	GROUP Unknown

EXAMINER INITIAL	OTHER DOCUMENTS (INCLUDING AUTHOR, TITLE, DATE, PERTINENT PAGES, ETC.)	
Num	79	Wong, W., J. Dimitroulakos, et al. (2002). "HMG-CoA Reductase Inhibitors and the Malignant Cell: the Statin Family of Drugs as Triggers of Tumor-Specific Apoptosis." Leukemia 16: 508-519.
WM	80	Yu, W., et al., (1999). "Induction of Apoptosis in Human Breast Cancer Cells by Tocopherols and Tocotrienols," Nutrition and Cancer, 33(1):26-32.

EXAMINER

DATE CONSIDERED

EXAMINER: INITIAL IF CITATION CONSIDERED, WHETHER OR NOT CITATION IS IN CONFORMANCE WITH MPEP 609; DRAW LINE THROUGH CITATION IF NOT IN CONFORMANCE AND NOT CONSIDERED, INCLUDE COPY OF THIS FORM WITH NEXT COMMUNICATION TO APPLICANT.